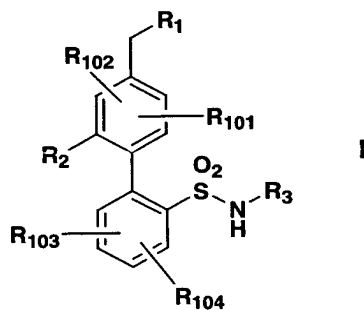


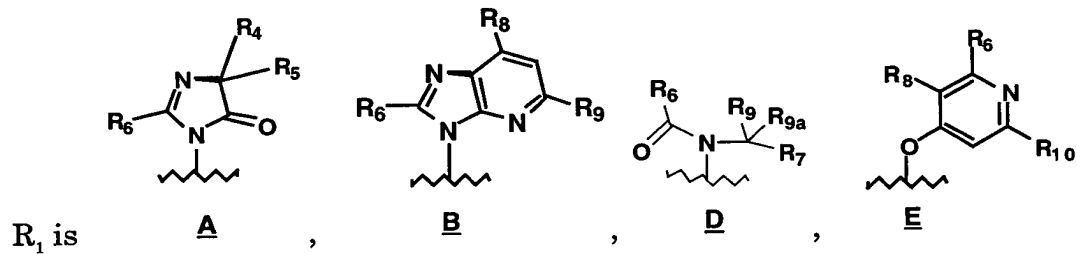
What is claimed is:

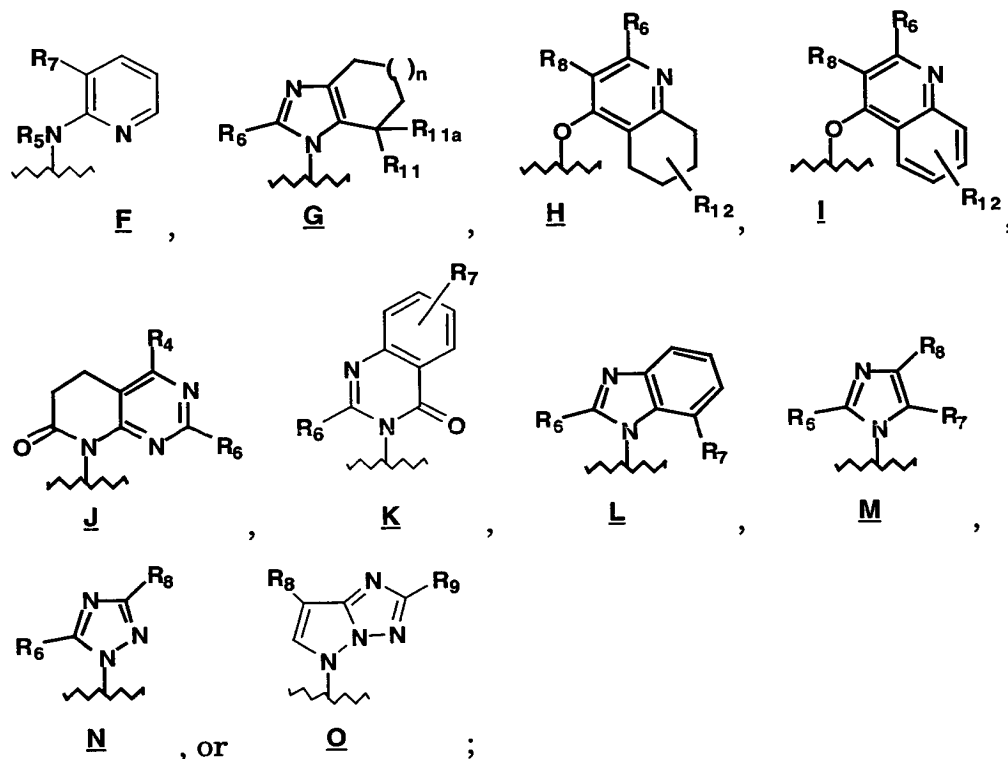
1. A compound of the following formula I, or an enantiomer, diastereomer, salt or metabolite thereof:

5



wherein:





5 R_2 is hydrogen, halogen, -CHO, alkyl, haloalkyl, (cycloalkyl)alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, aryloxy alkoxyalkoxy, cyano, hydroxy, hydroxyalkyl, nitro, $-\text{CH}(\text{OR}_{13})(\text{OR}_{14})$, $-(\text{CH}_2)_w\text{Y}$; with the proviso that when R_1 is B, R_2 is not hydrogen, halogen, alkyl, haloalkyl, alkoxy, hydroxyalkyl, nitro, $-(\text{CH}_2)_w\text{NR}_{19}\text{R}_{20}$ or $-\text{NH}\text{SO}_2\text{R}_{22}$;

10 R_3 is heteroaryl;

R_4 and R_5 are each independently alkyl, hydroxyalkyl, cycloalkyl, hydroxy substituted cycloalkyl, alkoxyalkyl, or hydroxy substituted alkoxyalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl, cyclohexyl, tetrahydrofuranyl or tetrahydropyranyl ring which may
 15 be optionally substituted with one or more hydroxy groups;

R_6 is alkyl, hydroxyalkyl, haloalkyl, hydroxy substituted haloalkyl, cycloalkyl, hydroxy substituted cycloalkyl, (cycloalkyl)alkyl, hydroxy substituted (cycloalkyl)alkyl, aralkyl, alkoxy, hydroxy substituted alkoxy, alkoxyalkyl, hydroxy substituted alkoxyalkyl, or
 20 $-\text{NR}_{16}\text{R}_{17}$;

R_7 is $-(CH_2)_w-CO_2R_{15}$, $-(CH_2)_w-(C=O)NR_{16}R_{17}$, $-(CH_2)_w-NR_{15}(C=O)NR_{16}R_{17}$, $-(CH_2)_w-CH_2OH$, $-(CH_2)_w-(C=O)R_{15}$, tetrazolyl, oxadiazolyl or triazolyl wherein said tetrazolyl, oxadiazolyl or triazolyl may optionally be substituted with hydrogen, alkyl, hydroxy or halogen;

5 R_8 , R_9 , R_{9a} , R_{10} and R_{12} are each independently hydrogen, halogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, aryl, heteroaryl, arylalkyl, alkylthioalkyl, alkoxy or alkoxyalkyl, or R_9 and R_{9a} together with the carbon atom to which they are bonded form a cycloalkyl ring;

10 R_{11} and R_{11a} are each independently hydrogen, alkoxy, or together form a carbonyl;

R_{13} and R_{14} are alkyl or together form a five to six-membered ring;

R_{15} , R_{16} and R_{17} are independently hydrogen, alkyl, hydroxyalkyl, cycloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, aralkyl, heterocycloalkyl,

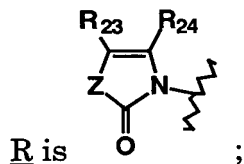
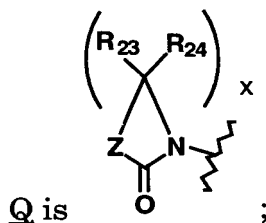
15 aryl, heteroaryl or $-(CH_2)_wQ$, or R_{16} and R_{17} may together form a four to six-membered heterocyclic ring;

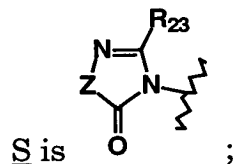
n is 1 or 2;

w is 0, 1, or 2;

Y is heteroaryl, $-COOH$, $-COOR_{18}$, $-CONR_{19}R_{20}$, $-NR_{19}R_{20}$, $-NR_{19}-OR_{20}$, -

20 $NR_{21}(C=O)R_{22}$, $-NR_{21}(C=O)NR_{19}R_{20}$, $-N(R_{19})-(alk)-NR_{21}(C=O)R_{22}$, $-NR_{21}(C=O)OR_{18}$, $-NR_{21}SO_2R_{22}$, $-SO_2R_{22}$, Q, R or S;

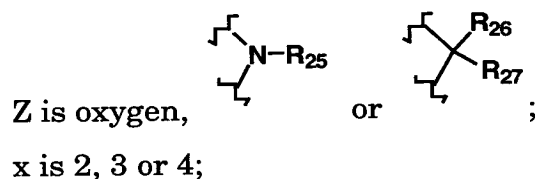




R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, haloalkyl, alkoxyalkyl, cycloalkyl, alkenyl, alkynyl, aryl, aralkyl, heteroaryl, or R_{19} and R_{20} may together form a four to seven-membered

5 heterocyclic ring;

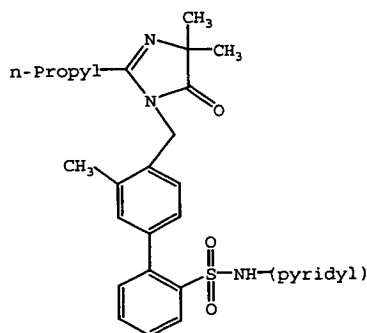
R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;



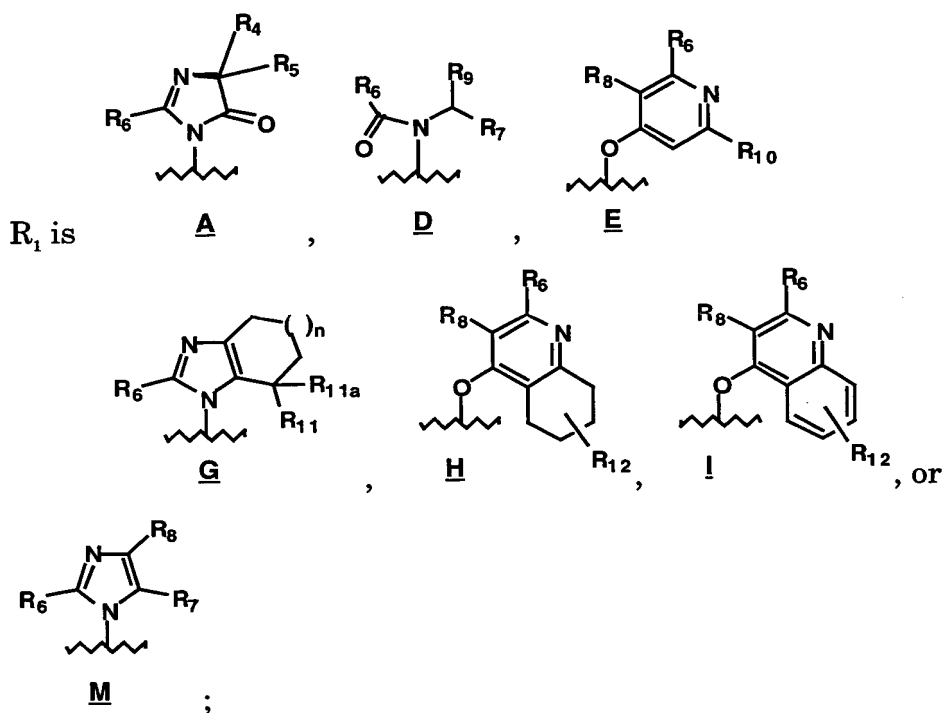
10 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

R_{101} , R_{102} , R_{103} and R_{104} are each independently hydrogen, halogen, -CHO, alkyl, haloalkyl, (cycloalkyl)alkyl, alkenyl, alkynyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, cyano, hydroxy, hydroxyalkyl, nitro, -CH(OR₁₃)(OR₁₄), or -(CH₂)_wY; wherein said rings; aryl alone or as part of another group; or heteroaryl alone or as part of another group may each optionally be substituted by one or more hydrogen, halogen, cyano, alkyl, hydroxyalkyl, alkoxy, nitro or trifluoromethyl groups;

20 provided that when R_1 is A said compound is other than



2. A compound of claim 1, wherein



R_2 is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$, or when R_1 is D , R_2 is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$;

R_3 is isoxazolyl, pyridizynyl, pyrazinyl or pyrimidinyl, each optionally independently substituted with one to three substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;

R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

R_6 is alkyl, haloalkyl, cycloalkyl or alkoxy;

R_7 is $-CO_2R_{15}$, $-(C=O)NR_{16}R_{17}$ or $-CH_2OH$;

R_8 , R_9 , R_{10} and R_{12} are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

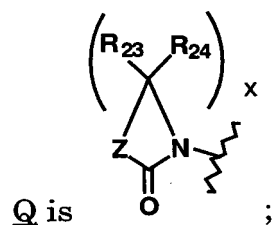
R_{11} and R_{11a} are each independently hydrogen, alkoxy, or together form a carbonyl;

R_{15} , R_{16} and R_{17} are independently hydrogen, alkyl or cycloalkyl or R_{16} and R_{17} may together form a four to six-membered heterocyclic ring;

5 n is 1 or 2;

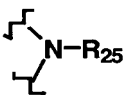
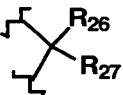
w is 0, 1, or 2;

Y is $-\text{COOR}_{18}$, $-\text{NR}_{21}(\text{C}=\text{O})\text{R}_{22}$, $-\text{NR}_{21}(\text{C}=\text{O})\text{NR}_{19}\text{R}_{20}$, $-\text{NR}_{21}(\text{C}=\text{O})\text{OR}_{18}$, $-\text{NR}_{21}\text{SO}_2\text{R}_{22}$, $-\text{SO}_2\text{R}_{22}$ or Q ;



10 R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four to seven-membered heterocyclic ring;

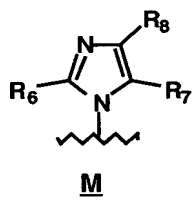
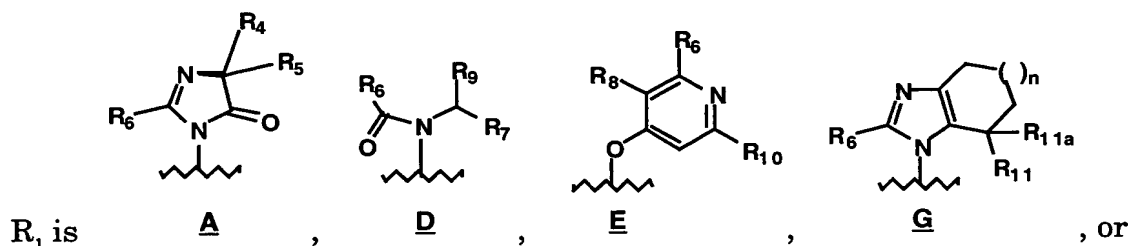
R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

15 Z is oxygen,  or  ;

x is 2, 3 or 4;
 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

20 R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, alkoxy or alkyl.

3. A compound of claim 1, wherein



R_2 is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, hydroxyalkyl, or $-(CH_2)_wY$, or when R_1 is D, R_2 is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$;

R_3 is isoxazolyl, optionally independently substituted with one or two substituents selected from hydrogen, halogen, cyano, alkyl, alkoxy, trifluoromethyl or nitro;

R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;

R_6 is alkyl, haloalkyl, cycloalkyl or alkoxy;

R_7 is $-\text{CO}_2R_{15}$ or $-(\text{C}=\text{O})\text{NR}_{16}R_{17}$;

R_8 , R_9 and R_{10} are each independently hydrogen, halogen, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;

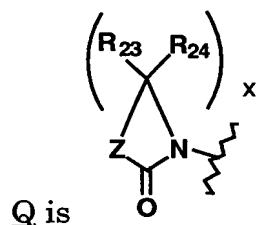
R_{11} and R_{11a} together form a carbonyl;

R_{15} , R_{16} and R_{17} are independently hydrogen, alkyl, or cycloalkyl or R_{16} and R_{17} may together form a four to six-membered heterocyclic ring;

n is 2;

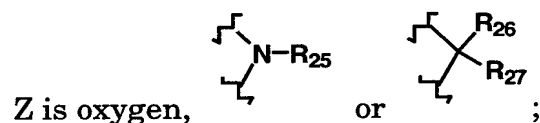
w is 0, 1, or 2;

Y is $-\text{NR}_{21}(\text{C}=\text{O})R_{22}$, $-\text{NR}_{21}(\text{C}=\text{O})\text{NR}_{19}R_{20}$, $-\text{NR}_{21}(\text{C}=\text{O})\text{OR}_{18}$, $-\text{NR}_{21}\text{SO}_2R_{22}$, $-\text{SO}_2R_{22}$ or Q;



R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four to seven-membered heterocyclic ring;

- 5 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;

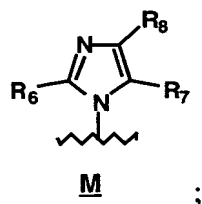
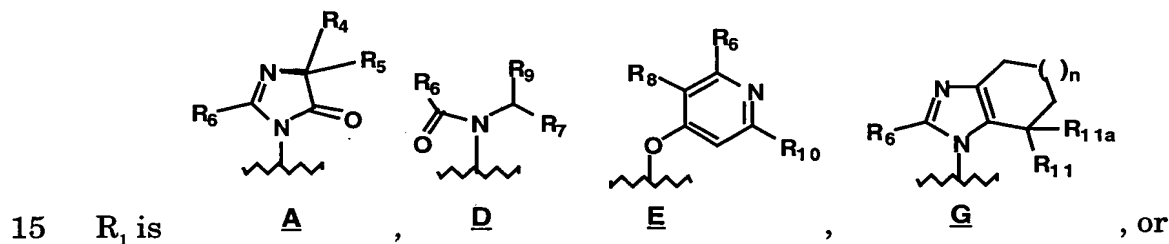


x is 2, 3 or 4;

- 10 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26} and R_{27} may together form a three to seven-membered cycloalkyl ring;

R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, or alkyl.

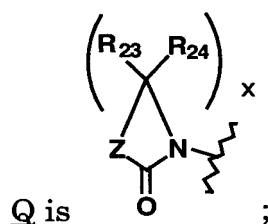
4. A compound of claim 1, wherein



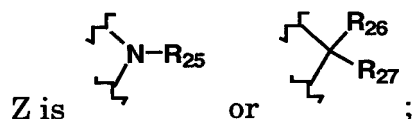
R_2 is alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$; or when R_1 is D, R_2

is hydrogen, alkyl, haloalkyl, (cycloalkyl)alkyl, alkoxyalkyl, haloalkoxyalkyl, alkoxy, alkoxyalkoxy, hydroxyalkyl, or $-(CH_2)_wY$;

- R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two
 5 substituents selected from alkyl or halogen;
 R_4 and R_5 are each independently alkyl, cycloalkyl, or R_4 and R_5 together form a cyclobutyl, cyclopentyl or cyclohexyl ring;
 R_6 is alkyl, haloalkyl, cycloalkyl or alkoxy;
 R_7 is $-(C=O)NR_{16}R_{17}$;
 10 R_8 , R_9 , and R_{10} are independently H, alkyl, cycloalkyl, alkoxy or alkoxyalkyl;
 n is 2;
 w is 0, 1, or 2;
 Y is $-NR_{21}(C=O)R_{22}$, $-NR_{21}(C=O)NR_{19}R_{20}$, $-NR_{21}(C=O)OR_{18}$, $-NR_{21}SO_2R_{22}$
 15 or Q ;



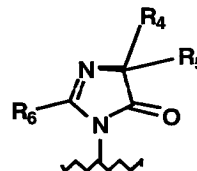
- R_{18} , R_{19} , R_{20} , R_{21} and R_{22} are each independently hydrogen, alkyl, cycloalkyl, or R_{19} and R_{20} may together form a four-, five-, six- or to seven-membered heterocyclic ring;
 20 R_{23} and R_{24} are each independently hydrogen, alkyl or cycloalkyl, or may together form a three to seven membered cycloalkyl ring;



- x is 2;
 R_{25} , R_{26} and R_{27} are each independently hydrogen, alkyl or cycloalkyl, or R_{26}
 25 and R_{27} may together form a three-, four-, five, six- or seven-membered cycloalkyl ring;

R_{101} , R_{102} , R_{103} , and R_{104} are each independently hydrogen, halogen, or alkyl.

5. A compound of claim 1, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

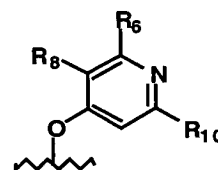


6. A compound of claim 5, wherein R_1 is A.

7. A compound of claim 6, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.

8. A compound of claim 6, wherein R_2 is $-CH_2Y$.

9. A compound of claim 8, wherein Y is Q.

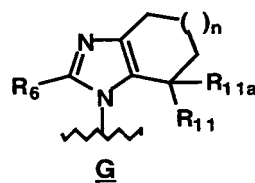


10. A compound of claim 5, wherein R_1 is E.

11. A compound of claim 10, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.

12. A compound of claim 10, wherein R_2 is $-CH_2Y$.

13. A compound of claim 12, wherein Y is Q.



14. A compound of claim 5, wherein R_1 is G.
15. A compound of claim 14, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.
16. A compound of claim 14, wherein R_2 is $-CH_2Y$.
- 10 17. A compound of claim 16, wherein Y is Q.
18. A compound of claim 1, wherein R_2 is alkoxyalkyl alkyl, haloalkyl or haloalkoxyalkyl
- 15 19. A compound of claim 18, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.
- 20 20. A compound of claim 1, wherein R_2 is $-CH_2Y$.
21. A compound of claim 20, wherein R_3 is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.
- 25 22. A compound of claim 20, wherein Y is Q.

23. A compound of claim 22, wherein R₃ is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen.

5 24. A compound of claim 1, wherein said compound is selected from the group consisting of

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(2-pyrazinyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-chloro-2-pyrazinyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3,6-dimethyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;

5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3-methoxy-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

(S)-N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;

15 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(2-oxo-1-pyrrolidinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

(S)-2-[N-[2'-[[N-(3-Methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

30 (S)-2-[N-[2'-[[N-(4-Bromo-3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]
-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-
N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-
(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-
(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-
10 (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
(methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
15 (carboxy)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-
biphenyl]-2-sulfonamide;
- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-
methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-
[1,1'-biphenyl]-2-sulfonamide;
- 20 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-
dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(1,3,5-trimethylpyrazol-4-yl)
[1,1'-biphenyl]-2-sulfonamide;
- 25 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(3-methylisoxazol-5-yl) [1,1'-
biphenyl]-2-sulfonamide;
- 4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-
dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-
30 (3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-N-(3-methoxy-5-methyl-2-pyrazinyl)[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-bromo-3-methyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylpropanamide;
- 20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylcyclopropanecarbox-amide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N,2-dimethylpropanamide;
- 25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-methylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 methoxy-N-methylacetamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-4-pentynamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylcyclobutanecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,3-dimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 15 N,2,2-trimethylpropanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 methoxy-N-methylpropanamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 ethoxy-N-methylacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-2-furancarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,4-dimethylpentanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 30 methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-3-thiophenecarboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylcyclopentaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylcyclohexanecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,3-dimethylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 15 methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 fluoro-N-methylbenzamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
 fluoro-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylcyclohexaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 30 fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
fluoro-N-methylbenzeneacetamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
fluoro-N-methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
N,N',N'-trimethylurea;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N'-
(1,1-dimethylethyl)-N-methylurea;

15 [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl)methyl)methylcarbamic acid ethyl ester;

[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl)methyl)methylcarbamic acid 2-methylpropyl ester;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
N,3,3-trimethylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
methyl-2-pyridinecarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
methyl-3-pyridinecarboxamide;

30 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
methyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,1-dimethyl-1H-pyrrole-2-carboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,5-dimethyl-2-pyrazinecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,3,5-trimethyl-4-isoxazolecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 15 N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,5-dimethyl-2-thiophenecarboxamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
 cyano-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 methoxy-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 30 chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 chloro-N-methylbenzamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
 chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 2,3-difluoro-N-methylbenzamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 3,4-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 15 3,5-difluoro-N-methylbenzamide;

4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-
 yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
 yl)methyl]-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 20 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 ethoxy-N-methyl-2-thiophenecarboxamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 25 dimethyl-5-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-
 sulfonamide;

N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 yl)methyl]-N-(1-oxopentyl)-L-valine methyl ester;

N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
 30 yl)methyl]-N-(1-oxopentyl)-L-valine;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidiny)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl)methyl]-N,3,3-trimethylbutanamide;

N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

10 N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;

4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

20 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

2'-(Cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 5 2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[[2,2,2-
- 10 trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]benzeneacetamide;
- 15 N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-dimethylbutanamide;
- 2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-isoleucinamide;
- N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
- 25 isoleucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
- 30

- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
- 5 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
- N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-leucinamide;
- 10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
- 15 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
- 20 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
- N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
- 25 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
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- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;
- 5 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;
- 10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-cyano[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 15 N,N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 20 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 25 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[[(5,6,7,8-tetrahydro-2-ethyl-4-quinolinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

10 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

15 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-yl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)-N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

20 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;

25 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;

N2-[[2-Chloro-2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

30 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(trifluoromethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-(Cyclobutylcarbonyl)-N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

1-[[2-Chloro-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-
5 biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid;
4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

(S)-N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-
10 biphenyl]-4-yl)methyl]-N-[2-methyl-1-[(4-methyl-1-piperazinyl)carbonyl]propyl]pentanamide;

(S)-N-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-piperidiny)carbonyl]propyl]pentanamide;

N-(3,3-Dimethylbutyl)- N2-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4-Chloro-1-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-
25 biphenyl]-4-yl)methyl]-2-propyl-1H-imidazole-5-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(1,2-dihydro-2-oxo-1-pyridinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-
30 biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-sulfonamide;
- 2-Butyl-4-chloro-1-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-1H-imidazole-5-carboxamide;
- 5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-methyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 10 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-ethyl-5,6,7,8-tetrahydro-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[2-propyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[[2-Ethyl-4-quinolinyloxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[5,6,7,8-tetrahydro-2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;
- 25 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester;
- 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester;
- 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-pyrrolidinyl)ethyl ester;
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- 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-pyrrolidinyl)propyl ester;
- 2'-Cyano-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;
- 2'-(Cyanomethyl)-N-(3,4-dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;
- 1-[[2-Chloro-2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;
- 3-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;
- 2-[[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]propylamino]-3-pyridinecarboxylic acid;
- 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2,7-Diethyl-5H-pyrazolo[1,5-b][1,2,4]triazol-5-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-[2-Butyl-3-[[2'-[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-methylethyl)urea;
- 2-[[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]propylamino]-N-methyl-3-pyridinecarboxamide;

- 4'-[[[(3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
 5 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
 10 3-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;
 4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
 20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
 25 N-[2-[[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl]methyl]methylamino]ethyl]acetamide;

4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
 [[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid
 ethyl ester;

N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
 5 4-yl)methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;

N2-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-
 4-yl)methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-L-
 valinamide;

2'-Chloro-N-(3,4-dimethyl-5-isoxazolyl)-4'-[[[(2-ethyl-4-
 10 quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2-ethyl-4-quinolinyloxy)methyl]-
 2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-
 N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 dimethyl-5-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-
 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 dimethyl-5-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-
 20 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 dimethyl-5-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2-
 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
 25 dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-
 sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-
 pyridinyl)oxy)methyl]-2'-ethoxymethyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
 30 dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3,3,3-trifluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3-fluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-difluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-trifluoroethyl)-N-(3,4-dimethylisoxazol-5-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3-methylisoxazol-5-yl)-2'-trifluoromethyl[1,1'-biphenyl]-2-sulfonamide;

N-(4-Bromo-3-methyl-5-isoxazolyl)-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-trifluoromethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-chloro-3-methyl-5-isoxazolyl)-2'-trifluoromethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-difluoroethoxymethyl)-N-(3,4-dimethylisoxazol-5-yl)[1,1'-biphenyl]-2-sulfonamide;

5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(2-methylpropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethylisoxazol-5-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

20

1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

25

1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

30

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-fluoroethoxymethyl) [1,1'-biphenyl]-2-

5 sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;

10 N-(3,4-Dimethyl-5-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[4-oxo-2-(3,3,3-trifluoropropyl)-1,3-diazaspiro[4.4]non-1-en-3-yl]methyl] [1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl) [1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

25 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

30 1-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

- N,4-Diethyl-1-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;
 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
 5 1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-carboxamide;
 1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
 10 N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
 N2-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxobutyl)-L-valinamide;
 15 N2-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxopentyl)-L-valinamide;
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
 4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
 20 N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide;
 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
 25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
 30

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;

5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy[1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-3'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-5-methoxy[1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy[1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-

20 sulfonamide;
4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-
25 N-(4,5-dimethyl-3-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)-4,5-dimethoxy[1,1'-biphenyl]-2-

30 sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4-methoxy-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;
- N²-(Cyclopropylcarbonyl)-N²-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-
- 15 valinamide;
- N²-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,3-dimethyl-N²-(1-oxobutyl)-L-valinamide;
- N²-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N²-(2-methyl-1-oxopropyl)-L-valinamide;
- 20 N²-(Cyclopentylcarbonyl)-N²-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
- N²-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N²-(1-oxobutyl)-L-valinamide;
- 25 N²-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(1-methylethyl)-N²-(1-oxobutyl)-L-valinamide;
- N²-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-(2-methoxyethyl)-N²-(1-oxobutyl)-L-valinamide;
- N-(Cyclopropylmethyl)-N²-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N²-(1-oxobutyl)-L-
- 30 valinamide;

N²-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N²-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;

N²-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N²-(1-oxopentyl)-L-valinamide;

5 N-Methyl-N²-[[2'-[[[(5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N²-(1-oxopentyl)-L-valinamide;

N²-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-ethyl-N²-(1-oxobutyl)-L-valinamide;

10 N²-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N²-(1-oxobutyl)-L-valinamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl]-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;

15 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 N-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(ethoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-1-[(1-oxopentyl)amino]cyclopentanamide;

and salts, enantiomers and diastereomer thereof.

25. A compound of claim 1, wherein said compound is selected from the group consisting of

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylamino)methyl][1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3-methyl-2-oxo-1-imidazolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-formyl-N-(3,4-dimethyl-5-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 20 (S)-N-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-[2-methyl-1-(3-methyl-1,2,4-oxadiazol-5-yl)propyl]pentanamide;
- N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 25 N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(2-methoxyethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[2-(ethoxymethyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
[(2-oxo-1-pyrrolidinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-
2-sulfonamide;

4'-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
5 [(3-methyl-2-oxo-1-imidazolidinyl)methyl]-N-(4,5-dimethyl-3-
isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

(S)-2-[N-[2'-[[N-(5-Methyl-3-isoxazolyl)amino]sulfonyl][1,1'-
biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

(S)-2-[N-[2'-[[N-(4-Bromo-5-methyl-3-isoxazolyl)amino]sulfonyl][1,1'-
10 biphenyl]-4-yl)methyl]-N-(1-oxopentyl)amino]-3,N-dimethylbutanamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]
-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(7-Methoxycarbonyl-2-ethoxybenzimidazol-1-yl)methyl]-
N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(7-Carboxy-2-ethoxybenzimidazol-1-yl)methyl]-N-
(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

4'-[(7-Methoxycarbonyl-2-ethylbenzimidazol-1-yl)methyl]-N-
(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

4'-[(7-Carboxy-2-ethylbenzimidazol-1-yl)methyl]-N-
20 (4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
(methoxycarbonyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-
isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;

2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-
25 (carboxy)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-
biphenyl]-2-sulfonamide;

2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N-
methylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-
[1,1'-biphenyl]-2-sulfonamide;

- 2'-[(3,3-Dimethyl-2-oxopyrrolidin-1-yl)methyl]-4'-[(2-ethoxy-7-(N,N-dimethylcarbamoyl)benzimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Ethylquinolin-4-yl)oxymethyl]-N-(5-methylisoxazol-3-yl)[1,1'-
5 biphenyl]-2-sulfonamide;
- 4'-[(5-Acetyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(5-Methoxycarbonyl-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(5-(N,N-dimethylcarbamoyl)-2-n-propyl-4-chloroimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-hydroxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
15 dimethyl-3-isoxazolyl)-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4-bromo-5-methyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-
20 pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(formylmethylamino)-methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
25 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylpropanamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylcyclopropanecarbox-amide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,2-dimethylpropanamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylbutanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 methoxy-N-methylacetamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-4-pentynamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 15 methylcyclobutanecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,3-dimethylbutanamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,2,2-trimethylpropanamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 methoxy-N-methylpropanamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 ethoxy-N-methylacetamide;

30 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-2-furancarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,4-dimethylpentanamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methyl-3-thiophenecarboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylcyclopentaneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 15 methylcyclohexanecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
 N,3-dimethylbenzamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
 methylbenzeneacetamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
 fluoro-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
 fluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
 30 fluoro-N-methylbenzamide;

- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
methylcyclohexaneacetamide;
- 5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-
fluoro-N-methylbenzeneacetamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-
fluoro-N-methylbenzeneacetamide;
- 10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-
fluoro-N-methylbenzeneacetamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
15 N,N',N'-trimethylurea;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N'-
(1,1-dimethylethyl)-N-methylurea;
- 20 [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl)methyl]methylcarbamic acid ethyl ester;
- [[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
yl)methyl]methylcarbamic acid 2-methylpropyl ester;
- 25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-
N,3,3-trimethylbutanamide;
- N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-
30 methyl-2-pyridinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methyl-3-pyridinecarboxamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,1-dimethyl-1H-pyrrole-2-carboxamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-N-
methyl-1,2,3-thiadiazole-4-carboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
15 N,5-dimethyl-2-pyrazinecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,3,5-trimethyl-4-isoxazolecarboxamide;

20 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,3-dimethyl-2-thiophenecarboxamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-
N,5-dimethyl-2-thiophenecarboxamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-3-
cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-
[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]methyl]-4-
30 cyano-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-methoxy-N-methylbenzamide;

5 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2-chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-chloro-N-methylbenzamide;

10 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-4-chloro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-2,3-difluoro-N-methylbenzamide;

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N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3,4-difluoro-N-methylbenzamide;

N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3,5-difluoro-N-methylbenzamide;

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4-Acetyl-N-[[4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-N-methylbenzamide;

25 N-[[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl)methyl]-3-ethoxy-N-methyl-2-thiophenecarboxamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(propylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)-L-valine methyl ester;

N-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-oxopentyl)-L-valine;

10 N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(2-propyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl)methyl]-N,3,3-trimethylbutanamide;

15 N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[[2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;

4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

25 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxylic acid methyl ester;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

30 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

- 2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-cyano-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-methyl[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-[(2,2,2-trifluoroethyl)amino]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]benzeneacetamide;
- N-[4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-yl]-3,3-dimethylbutanamide;
- 2'-Amino-4'-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-nitro[1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopropyl)-L-isoleucinamide;
- N2-(Cyclopropylcarbonyl)-N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-isoleucinamide;

- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-isoleucinamide;
- 5 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-isoleucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-leucinamide;
- 10 N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-leucinamide;
- 15 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(phenylacetyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-leucinamide;
- 20 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxohexyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxobutyl)-L-leucinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxopropyl)-L-valinamide;
- 25 N2-(Cyclopropylcarbonyl)- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl- N2-(1-oxo-3-phenylpropyl)-L-valinamide;
- 30

- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(phenylacetyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(3-methyl-1-oxobutyl)-L-valinamide;
- 5 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxohexyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl- N2-(1-oxobutyl)-L-valinamide;
- 10 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-ethyl- N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-hexyl- N2-(1-oxopentyl)-L-valinamide;
- N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-cyano[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 15 N2-[[2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-hydroxymethyl[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 4-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- 20 N,N-Dimethyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-carboxamide;
- N-Methyl-4-[(2-butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[[[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-
- 25 carboxamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-
- 30 pyridinyl)oxy]methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-methyl[1,1'-biphenyl]-2-sulfonamide;
- 5 N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 10 2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[5,6,7,8-tetrahydro-2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-fluoro[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(phenoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 20 N2-(Cyclopropylcarbonyl)-N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;
- N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;
- 25 N2-(Cyclopropylcarbonyl)- N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-L-valinamide;
- N2-[[2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(methoxymethyl)[1,1'-biphenyl]-4-yl]methyl]-N,N-dimethyl-N2-(1-oxobutyl)-L-valinamide;
- 30 N2-[[2-Chloro-2'-[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-(trifluoromethyl)[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;

5 N2-(Cyclobutylcarbonyl)-N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

1-[[2-Chloro-2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxylic acid;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylsulfonyl)amino][1,1'-biphenyl]-2-sulfonamide;

(S)-N-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[2-methyl-1-[(4-methyl-1-piperazinyl)carbonyl]propyl]pentanamide;

15 (S)-N-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-piperidiny)carbonyl]propyl]pentanamide;

N-(3,3-Dimethylbutyl)- N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-L-valinamide;

20 N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-[(4-fluorophenyl)methyl]-N2-(1-oxopentyl)-L-valinamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(methylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

25 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(propoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4-Chloro-1-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-propyl-1H-imidazole-5-carboxamide;

30 N-(4,5-dimethyl-3-isoxazolyl)-2'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-[(1,2-dihydro-2-oxo-1-pyridinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1H-pyrazol-1-ylmethyl)[1,1'-biphenyl]-2-sulfonamide;

2-Butyl-4-chloro-1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-1H-imidazole-5-carboxamide;

10 N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-methyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;

15 N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-ethyl-5,6,7,8-tetrahydro-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(2-propyl-4-quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(5,6,7,8-tetrahydro-2,4-dimethyl-7-oxopyrido[2,3-d]pyrimidin-8-yl)methyl][1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Ethyl-4-quinolinyl)oxy]methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(5,6,7,8-tetrahydro-2-ethyl-4-

25 quinolinyl)oxy]methyl [1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-N-methyl-3H-benzimidazole-4-carboxamide;

1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid phenylmethyl ester;

30 1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-phenylethyl ester;

1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 2-(2-oxo-1-pyrrolidinyl)ethyl ester;

5 1-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-1H-benzimidazole-7-carboxylic acid 3-(2-oxo-1-pyrrolidinyl)propyl ester;

2'-Cyano-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy)methyl][1,1'-biphenyl]-2-sulfonamide;

10 2'-(Cyanomethyl)-N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-N-methyl-2-propyl-3H-imidazole-4-carboxamide;

15 1-[[2-Chloro-2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-5-ethyl-2-propyl-3H-imidazole-4-carboxamide;

3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N-methyl-3H-benzimidazole-4-carboxamide;

20 3-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethoxy-N,N-dimethyl-3H-benzimidazole-4-carboxamide;

2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]propylamino]-3-pyridinecarboxylic acid;

25 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2,7-Diethyl-5H-pyrazolo[1,5-b][1,2,4]triazol-5-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

- N-[2-Butyl-3-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-3,4-dihydro-4-oxo-6-quinazolinyl]-N'-methyl-N'-(1-methylethyl)urea;
- 2-[[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]propylamino]-N-methyl-3-pyridinecarboxamide;
- 4'-[[[3-Methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl]-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(methoxymethyl)[1,1'-biphenyl]-2-sulfonamide;
- 3-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-2-ethyl-5-methyl-3H-imidazole-4-carboxamide;
- 4'-[(2-Butyl-3,4-dihydro-4-oxo-3-quinazolinyl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- N2-[[2'-[[[4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-2-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N2-(1-oxopentyl)-L-valinamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(4,4-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;
- 4'-[(3,5-Dibutyl-1H-1,2,4-triazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-[(3,3-dimethyl-2-oxo-1-pyrrolidinyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-[2-[[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl]-4-[(2-ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl][1,1'-biphenyl]-2-yl)methyl]methylamino]ethyl]acetamide;

4-[(2-Ethyl-5,7-dimethyl-3H-imidazo[4,5-b]pyridin-3-yl)methyl]-2'-
5 [[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-2-acetic acid
ethyl ester;

N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N2-(1-oxopentyl)-N-propyl-L-valinamide;

N2-[[2'-[(4,5-dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-
10 yl)methyl]-N2-(1-oxopentyl)-N-[(tetrahydro-2-furanyl)methyl]-L-
valinamide;

2'-Chloro-N-(4,5-dimethyl-3-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-dimethyl-3-isoxazolyl)-4'-[[2-ethyl-4-quinolinyloxy]methyl]-
15 2'-(trifluoromethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-chloro-
N-(4,5-dimethyl-3-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-[(2-methylpropoxy)methyl][1,1'-biphenyl]-2-
20 sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-[(ethylsulfonyl)amino][1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
25 dimethyl-3-isoxazolyl)-2'-[(2,2,2-trifluoroethoxy)methyl][1,1'-biphenyl]-2-
sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-
dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl][1,1'-biphenyl]-2-
sulfonamide;

30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-
pyridinyl]oxy]methyl]-2'-ethoxymethyl[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(3,3,3-trifluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(3-fluoropropyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(1,1-difluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2,2-trifluoroethyl)-N-(4,5-dimethylisoxazol-3-yl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-[(2-methyl)propoxy] [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-methoxyethoxy) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-butyl-N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(N-methoxy-N-methylaminomethyl) [1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-2'-(2,2-difluoroethoxymethyl)-N-(4,5-dimethylisoxazol-3-yl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-fluoroethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(3-methylbutyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(2-methypropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3-Difluorobutyl)-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethylisoxazol-3-yl)-2'-[(1,1-dimethylethoxy)methyl][1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methoxymethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

N,4-Diethyl-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-methyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-(1-methylethyl)-2-propyl-1H-imidazole-5-carboxamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethoxymethyl-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(5-Acetyl-4-ethyl-2-propylimidazol-1-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-methyl[1,1'-biphenyl]-2-sulfonamide;

- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
- 15 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-
- 20 N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
- 25 dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-5-methoxy[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)-4,5-dimethoxy[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-5'-chloro-N-(3,4-dimethyl-5-isoxazolyl)-2'-fluoro[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-3'-chloro-N-(3,4-dimethyl-5-isoxazolyl)[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4-methoxy-3'-methyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-fluoro-4-methoxy[1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-3'-methyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-5-fluoro[1,1'-biphenyl]-2-sulfonamide;

N²-(Cyclopropylcarbonyl)-N²-[[2'-[[[(3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-L-

25 valinamide;

N²-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N,3-dimethyl-N²-(1-oxobutyl)-L-valinamide;

N²-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N²-(2-methyl-1-oxopropyl)-L-valinamide;

N²-(Cyclopentylcarbonyl)-N²-[[2'-[[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-L-valinamide;

5 N²-[[2'-[[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl]-3-fluoro[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N²-(1-oxobutyl)-L-valinamide;

N²-[[2'-[[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(1-methylethyl)-N²-(1-oxobutyl)-L-valinamide;

N²-[[2'-[[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-(2-methoxyethyl)-N²-(1-oxobutyl)-L-valinamide;

10 N-(Cyclopropylmethyl)-N²-[[2'-[[[3,4-dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N²-(1-oxobutyl)-L-valinamide;

N²-[[2'-[[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N²-(1-oxobutyl)-N-(3-pyridinyl)-L-valinamide;

15 N²-[[2'-[[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-methyl-N²-(1-oxopentyl)-L-valinamide;

N-Methyl-N²-[[2'-[[[3-methyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N²-(1-oxopentyl)-L-valinamide;

20 N²-[[2'-[[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl)methyl]-N-ethyl-N²-(1-oxobutyl)-L-valinamide;

N²-[[2'-[[[3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-5'-fluoro[1,1'-biphenyl]-4-yl)methyl]-N-methyl-N²-(1-oxobutyl)-L-valinamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

25 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[2,6-dimethyl-3-methoxy-4-pyridinyl)oxy]methyl]-3'-fluoro[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

30 N-(3,4-Dimethyl-5-isoxazolyl)-3'-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-5-fluoro-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(hydroxymethyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-[[2'-[[[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl]-2-(ethoxymethyl)[1,1'-biphenyl]-4-yl)methyl]-1-[(1-oxopentyl)amino]cyclopentanamide;

and salts, enantiomers and diastereomer thereof.

10 26. A method for the treatment of an endothelin-dependent or angiotensin II-dependent disorder, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

15 27. A method for the treatment of hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

20 28. A method for the treatment of pulmonary hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25 29. A method for the treatment of primary pulmonary hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

30 30. A method for the treatment of low renin hypertension, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

31. A method for the treatment of male erectile dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

5 32. A method for the treatment of male or female sexual dysfunction, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

10 33. A method for the treatment of heart failure, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

15 34. A method for the treatment of atherosclerosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

20 35. A method for the treatment of restenosis, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

 36. A method for the treatment of endotoxemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25 37. A method for the inhibition of cell growth, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

30 38. A method for the treatment of cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

39. A method for the treatment of migraine, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

5

40. A method for the treatment of asthma, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

10

41. A method for the treatment of ischemia, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

42. A method for the treatment of subarachnoid hemorrhage,
15 comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

43. A method for the treatment of benign prostatic hypertrophy,
comprising the step of administering to a subject in need thereof an
20 amount effective therefor of at least one compound of claim 1.

44. A method for the treatment of renal, glomerular or mesangial
cell disorders, comprising the step of administering to a subject in need
thereof an amount effective therefor of at least one compound of claim 1.

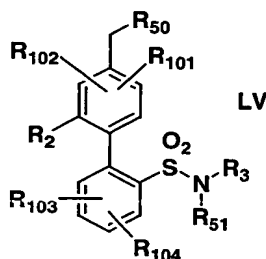
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45. A method for the treatment of acute or chronic renal failure,
comprising the step of administering to a subject in need thereof an
amount effective therefor of at least one compound of claim 1.

30 46. A pharmaceutical composition for the treatment of an
endothelin-dependent or angiotensin II-dependent disorder, comprising a

pharmaceutically acceptable vehicle or diluent and at least one compound of claim 1 in an amount effective therefor.

47. A compound of the formula



5

wherein

R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1;

R_{50} is hydroxy, chloro, bromo, iodo, $-\text{OSO}_2\text{-alkyl}$, or $-\text{OSO}_2\text{-aryl}$; and

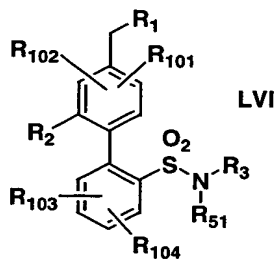
R_{51} is hydrogen, or a suitable nitrogen protecting group.

10

48. The compound of claim 47, wherein R_{51} is

$-\text{CH}_2\text{OCH}_2\text{CH}_2\text{OCH}_3$, $-\text{CH}_2\text{OCH}_2\text{CH}_2\text{Si}(\text{CH}_3)_3$, $-\text{CH}_2\text{OCH}_3$, or $-\text{CH}_2\text{OCH}_2\text{-aryl}$.

49. A compound of the formula



15

wherein R_1 , R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1; and

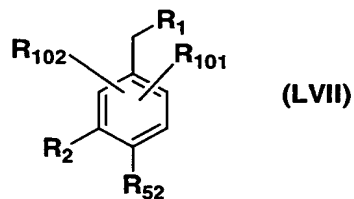
R_{51} is a suitable nitrogen protecting group.

50. The compound of claim 49, wherein R_{51} is $-\text{CH}_2\text{OCH}_2\text{CH}_2\text{OCH}_3$, -

20

$\text{CH}_2\text{OCH}_2\text{CH}_2\text{Si}(\text{CH}_3)_3$, $-\text{CH}_2\text{OCH}_3$, or $-\text{CH}_2\text{OCH}_2\text{-aryl}$.

51. A compound of the formula



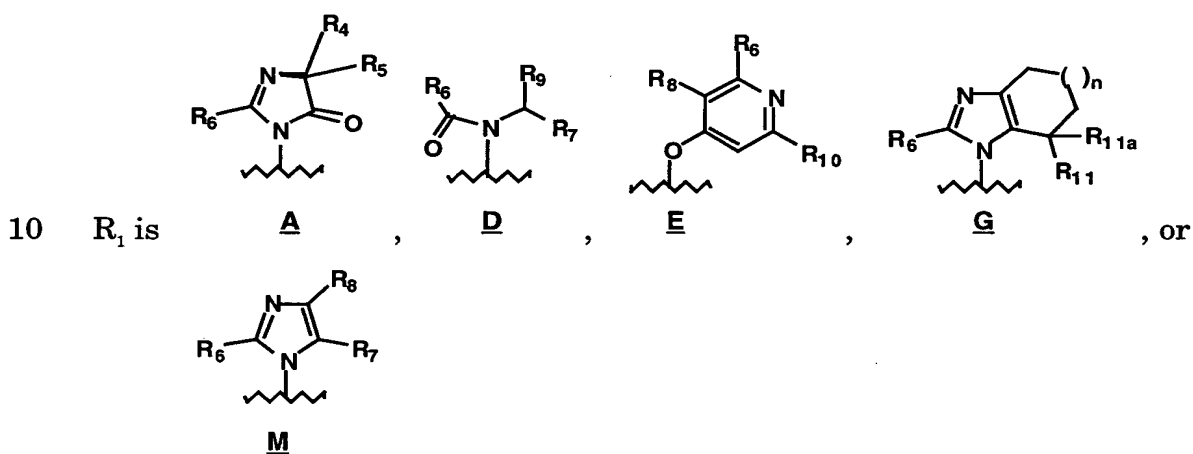
wherein

R_1 , R_2 , R_{101} and R_{102} are as defined in claim 1; and

5 R_{52} is chloro, bromo, iodo, or $-\text{OSO}_2\text{CF}_3$.

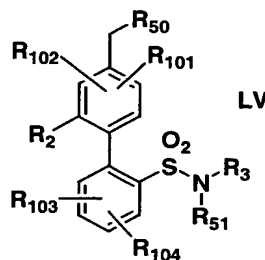
52. A compound of claim 51

wherein



53. A process for the preparation of a compound of claim 1, comprising at least one of the following steps:

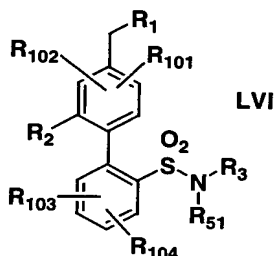
15 a) displacement of a leaving group R_{50} via the anion of a compound $R_1\text{-H}$ from a compound of the formula



wherein R_1 , R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, R_{50} is hydroxy, chloro, bromo, iodo, $-\text{OSO}_2$ -alkyl or $-\text{OSO}_2$ -aryl, and R_{51} is hydrogen or a suitable nitrogen protecting group, using a Mitsunobu reaction or S_N1 or S_N2 displacement reaction, with removal of said nitrogen

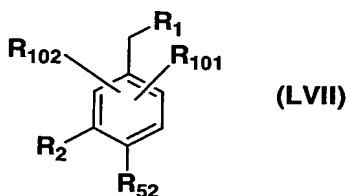
5 protecting group as appropriate;

b) removal of the nitrogen protecting group R_{51} from a compound of formula

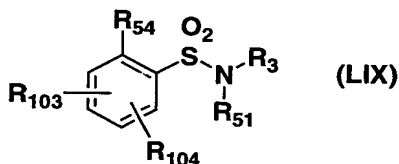


10 wherein R_1 , R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is a suitable nitrogen protecting group;

c) organometallic coupling of a compound of formula



15 with a compound of formula



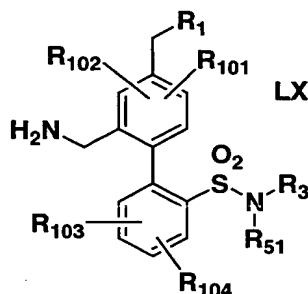
wherein

R_1 , R_2 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined for a compound of formula I and R_{51} is a suitable nitrogen protecting group;

20 R_{52} is chloro, bromo, iodo or $-\text{OSO}_2\text{CF}_3$ and R_{54} is a boronic acid, boronic ester or stannane derivative;

or R_{52} is a boronic acid, boronic ester or stannane derivative and R_{54} is chloro, bromo, iodo or $-\text{OSO}_2\text{CF}_3$;

d) acylation of a compound of the formula

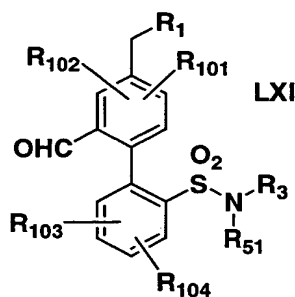


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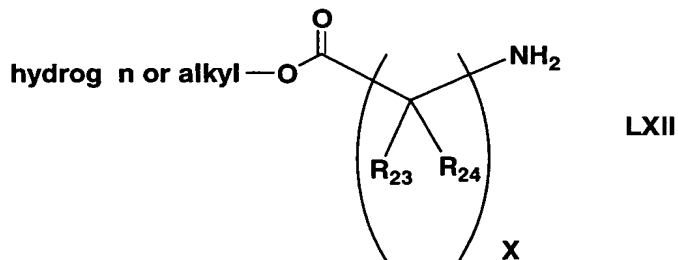
wherein R_1 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is hydrogen or a suitable nitrogen protecting group, with an acylating agent of the formula $R_{55}-(\text{C}=\text{O})R_{22}$, $R_{19}\text{-N}=\text{C}=\text{O}$, $R_{55}\text{-CO}_2R_{18}$, $R_{55}\text{-SO}_2R_{22}$, wherein R_{18} , R_{19} and R_{22} are as defined in claim 1 and R_{55} is an activating group for an acid, or made using an acid activating agent, with removal of said nitrogen protecting group as appropriate; or

10

e) reductive amination of a compound of formula



15 wherein R_1 , R_3 , R_{101} , R_{102} , R_{103} , and R_{104} are as defined in claim 1, and R_{51} is hydrogen or a suitable nitrogen protecting group, with an amine of the formula



wherein R_{23} , R_{24} , and x are as defined in claim 1, with removal of said nitrogen protecting group as appropriate.

5

54. A compound of claim 1, wherein said compound is selected from the group consisting of:

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;

15 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-ethyl[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)[1,1'-biphenyl]-2-sulfonamide;

20 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-ethyl-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

25 N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[[3-methoxy-2,6-dimethyl-4-pyridinyl]oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'-
 5 [[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-
 10 sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]- N-(3,4-dimethyl-5-isoxazolyl)-4'-
 15 [[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-4-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-
 20 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-4'-
 25 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethyl-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2,2-dimethylpropyl)-4'-
 30 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-2'-(2-ethoxyethyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide;

2'-[(1,1-Dimethylethoxy)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-4'-
5 [(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl][1,1'-biphenyl]-2-sulfonamide

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-
10 dimethyl-5-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

15 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-
25 sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofur-1-yl)[1,1'-
30 biphenyl]-2-sulfonamide;

- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
- 5 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-biphenyl]-2-sulfonamide;
- 10 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxyethyl)[1,1'-biphenyl]-2-sulfonamide;
- 20 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1-hydroxy-1-methylethyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,2,3,4-tetrahydrofuran-1-yl)[1,1'-biphenyl]-2-sulfonamide;
- 25 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;
- 30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(3,3,3-trifluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

10 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

15 N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

20 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

25 N-(4,5-Dimethyl-3-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)- 4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(2-fluoroethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

30 4'-[[2-(2,2-Difluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(2,2-Difluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

5 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

15 4'-[[2-(3,3,3-Trifluorobutyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

20 4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(ethoxymethyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

25 4'-[[2-(3,3,3-Trifluoropropyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-propyl[1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[(1,4,5,6,7,8-hexahydro-8-oxo-2-propyl-1-cycloheptimidazolyl)methyl]-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

10 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(3,4-dimethyl-5-isoxazolyl)-2'-(1,1-difluoropropyl)[1,1'-biphenyl]-2-sulfonamide;

1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

15 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

20 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

25 1-[[2'-[(3,4-Dimethyl-5-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;

1-[[2'-[(4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;

- 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- 5 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-N-methyl-2-propyl-1H-imidazole-5-carboxamide;
- 10 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-ethyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-propyl][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 15 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(2-fluoroethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 1-[[2'-[[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl-2-(ethoxymethyl)][1,1'-biphenyl]-4-yl]methyl]-4-ethyl-2-propyl-1H-imidazole-5-carboxamide;
- 20 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[6-ethyl-3-methoxy-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 25 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide;

- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-methoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 5 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 10 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 15 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-6-ethyl-2-methyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;
- 20 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;
- 25 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(3-ethoxy-2-methyl-6-propyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;
- 30 N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethyl [1,1'-biphenyl]-2-sulfonamide;

N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;

5 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-ethoxymethyl [1,1'-biphenyl]-2-sulfonamide;

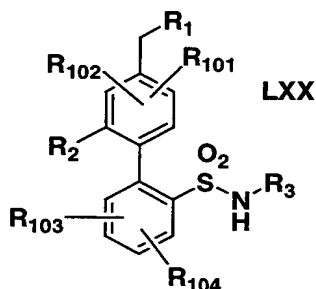
N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

10 N-(3,4-Dimethyl-5-isoxazolyl)-4'-[[[(2,6-dimethyl-3-ethoxy-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide;

and salts, enantiomers and diastereomers thereof.

55. A compound of formula LXX, or an enantiomer, diastereomer or salt thereof

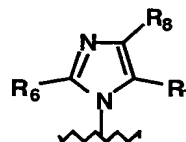
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wherein R₂, R₁₀₁, R₁₀₂, R₁₀₃, R₁₀₄ are as defined in claim 1;

R₃ is isoxazol-5-yl or isoxazol-3-yl independently substituted with two substituents selected from alkyl or halogen; and

20 R₁ is any group such that the resulting compound demonstrates affinity (IC₅₀) for both the AT₁ receptor and ET_A receptor of less than 5 micromolar at both receptors.



56. A compound of claim 5, wherein R₁ is

M

57. A compound of claim 56, wherein R_2 is alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R_{101} , R_{102} , R_{103} , R_{104} are each independently hydrogen, halogen, or alkyl.

5

58. A compound of claim 56, wherein R_2 is $-CH_2Y$.

59. A compound of claim 58, wherein Y is Q.

60. A crystalline form of 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide having a melting point of about 148°C.

10

61. N2-[[2'-[[4,5-Dimethyl-3-isoxazolyl)amino]sulfonyl][1,1'-biphenyl]-4-yl]methyl]-N-methyl-N2-(1-oxobutyl)-L-valinamide or a salt, enantiomer or diastereomer thereof.

15

62. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

20

63. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(2-fluoroethoxymethyl) [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

64. 4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-propyl [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

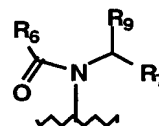
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65. N-(4,5-Dimethyl-3-isoxazolyl)-2'-ethoxymethyl-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

30

66. N-(4,5-Dimethyl-3-isoxazolyl)-2'-[(2-fluoroethoxy)methyl]-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl] [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.

5 67. N-(4,5-Dimethyl-3-isoxazolyl)-4'-[[3-methoxy-2,6-dimethyl-4-pyridinyl)oxy]methyl]-2'-propyl [1,1'-biphenyl]-2-sulfonamide or a salt, enantiomer or diastereomer thereof.



10 68. A compound of claim 5, where in R₁ is D.

69. A compound of claim 68, wherein R₂ is hydrogen, alkyl, haloalkyl, alkoxyalkyl or haloalkoxyalkyl and R₁₀₁, R₁₀₂, R₁₀₃, R₁₀₄ are each independently hydrogen, halogen, or alkyl.

15 70. A compound of claim 68 wherein R₂ is -CH₂Y.

71. A compound of claim 70, wherein Y is Q.

20 72. A method for the treatment of chronic obstructive pulmonary disease, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

25 73. A method for the alleviation of pain associated with prostate cancer, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

74. A method for preventing or reducing organ damage associated with the cell poliferative effects of endothelin, comprising the step of

administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

75. A method for reducing general morbidity and mortality
5 associated with endothelin-dependent or angiotensin II-dependent disorders, comprising the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

76. A method for the treatment of diabetic nephropathy, comprising
10 the step of administering to a subject in need thereof an amount effective therefor of at least one compound of claim 1.

77. A method for the treatment of dementia, comprising the step of administering to a subject in need thereof an amount effective therefor of
15 at least one compound of claim 1.

78. A compound of claim 1 wherein R_3 is other than pyridyl when R_1 is A.

79. The method of claim 26 wherein said compound of claim 1 is
20 administered prior to, simultaneously with or following the administration of at least one ACE inhibitor.

80. The method of claim 79 wherein said ACE inhibitor is
25 selected from captopril, zofenopril, fosinopril, ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril, or lisinopril.

81. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration
30 of at least one vasopectidase inhibitor.

82. The method of claim 81 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.

5 83. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one HMG CoA reductase inhibitor.

84. The method of claim 83 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin, atorvastatin,
10 simvastatin, NK-104 or ZD-4522.

85. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one anti-platelet agent.

15

86. The method of claim 85 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.

87. The method of claim 26 wherein said compound of claim 1 is
20 administered prior to, simultaneously with or following the administration of at least one anti-diabetic agent.

88. The method of claim 87 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.

25

89. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one beta-adrenergic agent.

90. The method of claim 89 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.

5 91. The method of claim 26 wherein said compound of claim 1 is administered prior to, simultaneously with or following the administration of at least one mineralocorticoid receptor antagonist.

10 92. The method of claim 91 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.

93. The pharmaceutical composition of claim 46 further comprising at least one ACE inhibitor.

15 94. The pharmaceutical composition of claim 93 wherein said ACE inhibitor is selected from captopril, zofenopril, fosinopril, ceranapril, alacepril, enalapril, delapril, pentopril, quinapril, ramipril, or lisinopril.

20 95. The pharmaceutical composition of claim 46 further comprising at least one vasopepsidase inhibitor.

96. The pharmaceutical composition of claim 95 wherein said vasopepsidase inhibitor is selected from omapatrilat or gemopatrilat.

25 97. The pharmaceutical composition of claim 46 further comprising at least one HMG CoA reductase inhibitor.

30 98. The pharmaceutical composition of claim 97 wherein said HMG CoA reductase inhibitor is selected from pravastatin, lovastatin, atorvastatin, simvastatin, NK-104 or ZD-4522.

99. The pharmaceutical composition of claim 46 further comprising at least one anti-platelet agent.

5 100. The pharmaceutical composition of claim 99 wherein said anti-platelet agent is selected from clopidigrel, ticlopidine, CS-747 or aspirin.

10 101. The pharmaceutical composition of claim 46 further comprising at least one anti-diabetic agent.

15 102. The pharmaceutical composition of claim 101 wherein said anti-diabetic agent is selected from biguanides or biguanide/glyburide combinations.

 103. The pharmaceutical composition of claim 46 further comprising at least one beta-adrenergic agent.

20 104. The pharmaceutical composition of claim 103 wherein said beta-adrenergic agent is selected from carvedilol or metoprolol.

 105. The pharmaceutical composition of claim 46 further comprising at least one mineralocorticoid receptor antagonist.

25 106. The pharmaceutical composition of claim 105 wherein said mineralocorticoid receptor antagonist is selected from spironolactone or eplerenone.

107. The compound of claim 62 that is (+)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.

5 108. The compound of claim 62 that is (-)-4'-[(2-Butyl-4-oxo-1,3-diazaspiro[4.4]non-1-en-3-yl)methyl]-N-(4,5-dimethyl-3-isoxazolyl)-2'-(ethoxymethyl) [1,1'-biphenyl]-2-sulfonamide.